

AMENDMENT

Kindly amend the application, without prejudice, without admission, without surrender of subject matter, and without any intention of creating any estoppel as to equivalents, as follows.

IN THE CLAIMS:

Kindly amend the claims, without prejudice, without admission, without surrender of subject matter, and without any intention of creating any estoppel as to equivalents, as follows:

96. (Currently Amended) A substantially pure, recombinant glycosylated erythropoietin, produced by a baculovirus expression system in cultured insect cells, wherein said erythropoietin has relative homogeneity or is purified to 95% or greater and said erythropoietin stimulates erythropoiesis and has an activity *in vivo* of at least 200,000 U/mg or of about 500,000 U/mg.

97. (Currently Amended) Erythropoietin of claim 96 wherein said erythropoietin stimulates erythropoiesis and has an activity *in vivo* of at least 200,000 U/mg.

98. (Currently Amended) Erythropoietin of claim 96 wherein said erythropoietin stimulates erythropoiesis and has an activity *in vivo* of at least 500,000 U/mg.

99. (Previously added) Erythropoietin of claim 96 produced by a method comprising:

culturing insect cells in at least one bioreactor whereby there is an insect cell culture,
wherein the insect cells contain a recombinant baculovirus containing exogenous DNA encoding erythropoietin,
supplying medium in at least one vessel whereby there is culture medium,
circulating culture medium and/or insect cell culture, whereby the bioreactor and vessel are in fluid communication and the insect cell culture and/or culture medium are in circulation,
delivering oxygen to the insect cell culture and/or culture medium, and
collecting the expressed product, and/or baculovirus and/or the cells.

100. (Previously added) Erythropoietin of claim 96 produced by a method comprising:

culturing insect cells in a bioreactor whereby there is an insect cell culture,
wherein the insect cells contain a recombinant baculovirus containing exogenous DNA encoding erythropoietin,

supplying culture medium in a vessel whereby there is culture medium,
circulating the insect cell culture through a dialysis means,
circulating culture medium through the dialysis means,
wherein the dialysis means in fluid communication with the bioreactor and the
vessel,
whereby

there is

a first, cell culture, loop between the bioreactor and the
dialysis means, and
a second, media replenishment, loop between the vessel
and the bioreactor,

performing dialysis between the culture medium and the cell culture, and
collecting the erythropoietin.

101. (Previously added) Erythropoietin as claimed in claim 100, wherein the
method further comprises:

delivering oxygen into the cell culture loop and measuring physical and/or chemical
parameter(s) of the cell culture and/or the culture medium.

102. (Previously added) Erythropoietin as claimed in claim 101, wherein the
method further comprises adjusting physical and/or chemical parameter(s) of the cell culture
and/or the culture medium in response to data from the measuring.

103. (Previously added) Erythropoietin as claimed in claim 101, wherein the
method further comprises measuring pH and measuring dissolved oxygen concentration,
adjusting physical and/or chemical parameter(s) of the cell culture and/or the culture medium in
response to data from the measuring, wherein the adjusting comprises adjusting temperature to
maintain a desired temperature, adjusting pH to maintain a desired pH, and adjusting dissolved
oxygen concentration and dissolved carbon dioxide concentrations, whereby the dissolved
carbon dioxide levels are adjusted in response to pH measurement(s).

104. (Previously added) Erythropoietin as claimed in claim 103, wherein the
method further comprises adjusting dissolved oxygen levels in response to dissolved oxygen
measurement(s), adjusting pH to a desired level in response to pH measurement(s) by adjusting

the dissolved carbon dioxide concentration such that dissolved carbon dioxide concentration is adjusted when pH varies from the desired level, and the dissolved oxygen measurement varies periodically as a function of time, adjusting the dissolved oxygen concentration so that the dissolved oxygen measurement varies from 30% to 90% or from 40% to 80% or from 50% to 70%; or, so that the dissolved oxygen measurement averages about 60%.

105 (Previously added) Erythropoietin as claimed in claim 104, wherein the adjusting of the dissolved oxygen concentration so that the dissolved oxygen measurement varies from 30% to 90%.

106 (Previously added) Erythropoietin as claimed in claim 104, wherein the adjusting of the dissolved oxygen concentration so that the dissolved oxygen measurement varies from 40% to 80%.

107 (Previously added) Erythropoietin as claimed in claim 104, wherein the adjusting of the dissolved oxygen concentration so that the dissolved oxygen measurement varies from 50% to 70%.

108 (Previously added) Erythropoietin as claimed in claim 104, wherein the adjusting of the dissolved oxygen concentration so that the dissolved oxygen measurement averages about 60%.

109. (Previously added) Erythropoietin as claimed in claim 104, wherein the method further comprises adjusting the dissolved oxygen concentration so that the dissolved oxygen measurement varies from high value to low value over about 10 to about 30 minutes or over about 20 minutes.

110. (Previously added) Erythropoietin as claimed in claim 103, wherein the method further comprises adjusting dissolved oxygen levels in response to dissolved oxygen measurement(s), and adjusting pH to a desired level in response to pH measurement(s) by adjusting the dissolved carbon dioxide concentration such that dissolved carbon dioxide concentration is adjusted when pH varies from the desired level, and the dissolved oxygen measurement varies periodically as a function of time, and wherein a plot of the dissolved oxygen measurement as a function of time comprises a sine wave.

111. (Previously added) Erythropoietin as claimed in claim 99 wherein the insect cells are *Spodoptera frugiperda* cells.

112. (Previously added) Erythropoietin as claimed in claim 100 wherein the insect cells are *Spodoptera frugiperda* cells.

113. (Previously added) Erythropoietin as claimed in claim 111 wherein the medium is serum free.

114. (Previously added) Erythropoietin as claimed in claim 112 wherein the medium is serum free.

115. (Previously added) Erythropoietin as claimed in claim 111 wherein the insect cells are *Spodoptera frugiperda* SF900+ cells.

116. (Previously added) Erythropoietin as claimed in claim 112 wherein the insect cells are *Spodoptera frugiperda* SF900+ cells.

117. (Withdrawn) An expressed product obtained by a method for growing cells comprising

culturing cells in at least one bioreactor whereby there is a cell culture,
supplying medium in at least one vessel whereby there is culture medium,
circulating culture medium and/or cell culture, whereby the bioreactor and vessel are in fluid communication and the cell culture and/or culture medium are in circulation, and delivering oxygen to the cell culture and/or culture medium,
wherein the cells contain a vector for replication of the vector and/or expression of exogenous nucleic acid molecules wherein the vector comprises a virus or a recombinant virus;
wherein the method further comprises collecting expressed product, and/or baculovirus and/or the cells.

118. (Withdrawn) An expressed product obtained by a method for growing cells comprising

culturing cells in a bioreactor whereby there is a cell culture, wherein the cells contain a vector for replication of the vector and/or expression of exogenous nucleic acid molecules wherein the vector comprises a virus or a recombinant virus,
supplying culture medium in a vessel where by there is culture medium,
circulating the cell culture through a dialysis means,
circulating culture medium through the dialysis means,
wherein the dialysis means in fluid communication with the bioreactor and the vessel,

whereby

there is

a first, cell culture, loop between the bioreactor and the dialysis means,
and

a second, media replenishment, loop between the vessel and the
bioreactor,

performing dialysis between the culture medium and the cell culture,
and collecting expressed product, and/or baculovirus and/or the cells.

119. (Withdrawn) The expressed product of claim 118, wherein the method by which
the product is obtained further comprises

delivering oxygen into the cell culture loop,

and measuring physical and/or chemical parameter(s) of the cell culture and/or the culture
medium.

120. (Withdrawn). The expressed product of claim 119, wherein the method by which
the product is obtained further comprises

adjusting physical and/or chemical parameter(s) of the cell culture and/or the culture
medium in response to data from the measuring.

121. (Withdrawn) The expressed product of claim 120, wherein the adjusting
comprises adjusting temperature to maintain a desired temperature, and wherein the adjusting
comprises adjusting dissolved oxygen concentration and adjusting dissolved carbon dioxide
concentration, whereby in response to pH measurement(s), dissolved carbon dioxide levels are
adjusted.

122. (Withdrawn) The expressed product of claim 121 wherein the adjusting includes
adjusting dissolved oxygen levels in response to dissolved oxygen measurement(s), wherein the
adjusting comprises adjusting pH to a desired level in response to pH measurement(s) by
adjusting the dissolved carbon dioxide concentration such that dissolved carbon dioxide
concentration is adjusted when pH varies from the desired level, and the dissolved oxygen
measurement varies periodically as a function of time, and wherein the adjusting includes
adjusting the dissolved oxygen concentration so that the dissolved oxygen measurement varies
from 30% to 90% or from 40% to 80% or from 50% to 70%; or, so that the dissolved oxygen
measurement averages about 60%.

123. (Withdrawn) The expressed product of claim 122, wherein the adjusting includes adjusting the dissolved oxygen concentration so that the dissolved oxygen measurement varies from high value to low value over about 10 to about 30 minutes or over about 20 minutes.

124. (Withdrawn) The expressed product of claim 121, wherein the adjusting includes adjusting dissolved oxygen levels in response to dissolved oxygen measurement(s), wherein the adjusting comprises adjusting pH to a desired level in response to pH measurement(s) by adjusting the dissolved carbon dioxide concentration such that dissolved carbon dioxide concentration is adjusted when pH varies from the desired level, and the dissolved oxygen measurement varies periodically as a function of time, and wherein a plot of the dissolved oxygen measurement as a function of time comprises a sin wave.